

1. A method of treating an autoimmune disease in a subject, the method comprising administering an effective amount of one or more of:

- a B-cell Activating Factor (BAFF) polypeptide;
- a BAFF polypeptide and an agent that promotes survival and/or migration of gut-derived commensal-reactive B cells to the central nervous system of the subject; or
- a BAFF polypeptide and a gut commensal that increases IgA levels to the subject.

2. The method of claim 1, wherein the autoimmune disease is a non-systemic organ-specific autoimmune disease.

3. The method of claim 1 or 2, wherein the autoimmune disease is multiple sclerosis.

4. The method of any one of claims 1-3, wherein the commensal-reactive B cells are IgA+ plasmablasts and/or plasma cells.

5. The method of any one of claims 1-4, wherein the commensal-reactive B cells express IL-10 and/or iNOS.

6. The method of any one of claims 1-5, wherein the agent that promotes survival and/or migration of gut-derived commensal-reactive B cells is a cytokine or a chemokine.

7. The method of any one of claims 1-5, wherein the agent that promotes survival and/or migration of gut-derived commensal-reactive B cells is IL-10 and/or iNOS.

8. The method of any one of claims 1-7, wherein the gut commensal is a commensal microbe.

9. The method of any one of claims 1-8, wherein the administering an effective amount of a gut commensal comprises oral or rectal administration of a microbe or community of microbes.

10. The method of any one of claims 1-9, wherein the gut commensal comprises *Tritrichomonas musculus* or a gut microbial community that has been modified by carriage of *Tritrichomonas musculus*.

11. A method of reducing inflammation in a subject, the method comprising administering an effective amount of one or more of:

- a B-cell Activating Factor (BAFF) polypeptide;
- a BAFF polypeptide and an agent that promotes survival and/or migration of gut-derived commensal-reactive B cells to the central nervous system of the subject; or
- a BAFF polypeptide and a gut commensal that increases IgA levels to the subject.

12. The method of claim 11, wherein the inflammation is reduced in the periphery of the subject.

13. The method of claim 11, wherein the inflammation is reduced in the central nervous system.

14. The method of any one of claims 11-13, wherein the inflammation is neuroinflammation.

15. The method of claim 14, wherein the neuroinflammation is caused by multiple sclerosis.

16. The method of any one of claims 11-15, wherein the commensal-reactive B cells are IgA+ plasmablasts and/or plasma cells.

17. The method of any one of claims 11-16, wherein the commensal-reactive B cells express IL-10 and/or iNOS.

18. The method of any one of claims 11-17, wherein the agent that promotes survival and/or migration of gut-derived commensal-reactive B cells is a cytokine or a chemokine.

19. The method of any one of claims 11-17, wherein the agent that promotes survival and/or migration of gut-derived commensal-reactive B cells is IL-10 and/or iNOS.

20. The method of any one of claims 11-19, wherein the gut commensal is a commensal microbe.

21. The method of any one of claims 11-20, wherein the administering an effective amount of a gut commensal comprises oral or rectal administration of a microbe or community of microbes.

22. The method of any one of claims 11-21, wherein the gut commensal comprises *Tritrichomonas musculus* or a gut microbial community that has been modified by carriage of *Tritrichomonas musculus*.

23. A method of enriching gut-derived commensal-reactive IgA+ plasmablasts and/or plasma cells in the central nervous system of a subject, the method comprising administering an effective amount of one or more of:

- a B-cell Activating Factor (BAFF) polypeptide;
- a BAFF polypeptide and an agent that promotes survival and/or migration of gut-derived commensal-reactive B cells to the central nervous system of the subject; or
- a BAFF polypeptide and a gut commensal that increases IgA levels to the subject.

24. The method of claim 23, wherein the gut-derived commensal-reactive IgA+ plasmablasts and/or plasma cells express IL-10 and/or iNOS.

25. The method of claim 23 or 24, wherein the subject has an autoimmune disease.

26. The method of any one of claims 23-25, wherein the subject has a non-systemic organ-specific autoimmune disease.

27. The method of claim any one of claims 23-36, wherein the subject has multiple sclerosis.

28. The method of any one of claims 23-27, wherein the agent that promotes survival and/or migration of gut-derived commensal-reactive B cells is a cytokine or a chemokine.

29. The method of any one of claims 23-27, wherein the agent that promotes survival and/or migration of gut-derived commensal-reactive B cells is IL-10 and/or iNOS.

30. The method of any one of claims 23-29, wherein the gut commensal is a commensal microbe.

31. The method of any one of claims 23-30, wherein the administering an effective amount of a gut commensal comprises oral or rectal administration of a microbe or community of microbes.

32. The method of any one of claims 23-31, wherein the gut commensal comprises *Tritrichomonas musculus* or a gut microbial community that has been modified by carriage of *Tritrichomonas musculus*.

33. A method of promoting survival of gut-derived commensal-reactive IgA+ plasmablasts and/or plasma cells in a subject to reduce inflammation in a tissue, the method comprising administering an effective amount of one or more of:

- a B-cell Activating Factor (BAFF) polypeptide;
- a BAFF polypeptide and an agent that promotes survival and/or migration of gut-derived commensal-reactive B cells to the central nervous system of the subject; or
- a BAFF polypeptide and a gut commensal that increases IgA levels to the subject.

34. The method of claim 33, wherein the gut-derived commensal-reactive IgA+ plasmablasts and/or plasma cells express IL-10 and/or iNOS.

35. The method of claim 33 or 34, wherein the subject has an autoimmune disease.